

Consulted
9/30/02

WO 00/37069

"NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC ACID
IN MEDICINE"

Inventor: 371 of PCT/Br99/00107 filed 12/17/1999
Technician: [Signature]

The present invention is relative to a compound made of alpha-hidroxi-propionic (or 2-hidroxi-propionic), (compound I), combined with 1,2,3-propanotriol pure (glicerine), or to the 1,2-propanodiol pure or serum, or to a balanced mixture of them or any other acceptable pharmaceutical vehicle, (compound II) to the attaining procedures of such a compound or to its utilization in Medicine.

Background of the Invention

The invention has got a compound consisted of the alpha-hidroxi-propionic acid (compound I), or to an acceptable pharmaceutical salt of the latter, or of an acceptable pharmaceutical solvato of the latter, or of an acceptable pharmaceutical catalyser of the latter, characterized by the following dilution of:

0,2 to 0,9ml, or 1,1 to 2,0ml, or 2,1 to 3,0ml, or 3,1 to 4,0 ml, or 4,1 to 5,0ml, or 5,1 to 6,0ml, or 6,1 to 10ml of compound I in 100ml of compound II; and 0,3 to 0,8 ml, or 0,4 to 0,7 ml, or 0,2 to 0,5ml, or 0,5 to 0,9 ml, or 1,1 to 1,5ml, or 1,5 to 2,0ml, or 2,1 to 3,0ml, or 3,1 to 4,0ml, or 4,1 to 5,0 ml or 5,1 to 10,0 ml of the active principle of compound I in 100ml of compound II.

Compound I, in one or more of the above-mentioned items combined with compound II is characterized by being suit to the intake in drops, via the nasal airways, or as a spraying solution, a spray, a microfine powder for insufflation or an acceptable pharmaceutical salt or an acceptable pharmaceutical solvate for the medicine addressed to the treatment of the highairways disturbances.

Summary of the Invention

There aren't in the medical and pharmaceutical literatures any statements about the active principle of compound I. On the other hand, there isn't an efficient medicine for the sinusitis treatment. What has been recorded in medical literature so far is the antibiotics massification which, besides its high cost, represents one of the biggest threats to the world public health, due to the development of resistant "cepas" (germs).

Detailed Description of the Invention

It's to be pointed out that the antibiotics massification leads only to the

09868793-100107

germs fight inside the organism or in its "doorway" when such disturbances are in acute crisis. During those crisis, the germs either in the nasal cavities or in the cheek bones located in external areas of the organism, in close contact with the external environment, aren't reached.

For a biologically active substance to carry out its duty, it's necessary to be positioned at the action location. The active principles are taken into the body through medicines. Therefore, it's necessary for them to be released in the location where the infectious agents are.

In fact, the antibiotic is a medicine for internal use and that's why it isn't efficient in the sinusitis treatment, taking into consideration that its release doesn't occur at the infection spot. As known, the sinusitis is an inflammation of the layer of the tissue that internally covers the cheek bones through little holes which communicate with the nasal cavity directly linked to the external environment.

As the application of compound I linked to compound II occurs at the nostrils, such a compound will work directly on the germs located in the nasal cavities and cheeks.

The first application effect in the nasal cavities and cheek bones of compound I linked to compound II is the "lisar" (dehydrating) of the germs that can be found there through its bactericide and bacteriostatic properties that are in contact .

After that, the hydrating and moistening effects of compound I linked to compound II, cause the increase in the nasal mucosa elasticity and its clearance. The action motion of the alpha-hidroxi-propionic acid keeps a more hemogeneous cornea layer, decreasing the superficial cellular cohesion. Those alpha-hidroxi-propionic acids promote a subtle exfoliation, leaving the nasal mucosa smoother and more homogeneous.

As mediate effects, there are also the modifications of the medium pH, facilitating the "Lactobacillus acidophyllus" and the "Bifidobacteria" growth. The Bifidobacteria are known for displaying inhibiting effects upon many other

09866793-100101

pathogenic germs, "in vitro" and "in vivo", such as "Candida albicans",
60 "Shighellas", "Clostridium", "Bacillus cereus", "Staphylococcus aureus", and
"Campylobacter jejuni", according to the researches of Aann and col. (1985), Tojo
and col. (1987), Tomoda and col. (1988).

It's known, as well, that the bifidobacteria in the large intestine
synthesize vitamins that are absorbed by the organism.

65 Bifidobacteria are still known for producing tiamine, riboflavine and
vitamins B6 and K. It's still proved that the bifidobacteria are able to synthetize the
complex B vitamins (Mutai, 1978).

In the cheek bones, the compound I linked to compound II changes the
medium pH, promoting the mucosa hydrating which will speed up the
70 bifidobacteria growth. The bifidobacteria, by competition, leaves out the
pathogenic bacteria found there, which are responsible for the cheek bones
infections. Then, the environmental adaptation to the new pH makes the cheek
bones prone to the bifidobacteria development, as it occurs in the intestines (Rassic
- 1989).

75 Well, similarly to the gastrointestinal tract, the respiratory system is open
to the external environment in order to facilitate the organism breathing. In fact,
the bifidobacteria and "Lactobacyllus acidophyllus" growth in the cheek bones, is
possible due to the optimum pH, determined by the active principles of compound
I linked to compound II.

80 Researches believe that the bifidobacteria, by competition, leave out the
large intestines putrefying bacteria which are responsible for the free radicals
release. The free radicals, being absorbed, will do the organism a lot of harm, such
as early aging. (Metchnikoff, 1938 and Linnus Pauling, 1965)

Therefore, the "Lactobacyllus acidophyllus" and Bifidobacteria
85 presences are beneficial to the cheek bones as well as to the intestines. One of the
Bifidobacteria effects as an effective pathogenic germ inhibitor is associated with
the production of lactats and acetats in small portions in the mechanism of

reaction in the chemical products resultant from the carbohydrates catabolism. Those elements and the pH inhibit the pathogenic bacteria growth. (Hughes, D.B., Hoover, D.G., BIFIDOBACTERIA, THEIR POTENTIAL FOR USE IN AMERICAN PRODUCTS).

The medicine utilization, represented by compound I linked to compound II is considered only by the otorhinolaryngologist clinics as a salutary alternative to the rhinitis and sinusitis treatment.

Carriers of such diseases feel considerable relief from the very first time they take the referred medicine.

The medicine, represented by compound I linked to compound II, has shown advantages upon any other medicine, for it isn't reabsorbed for being a product of cellular rejects.

At present, the sinusitis is treated with last generation antibiotics, not always with the desired results for not reaching the infection focus, which is inaccessible, and its massification leads to one of the biggest threats to the world public health due to the resistant "cepas" (germs) appearance _ what would justify this request at once.

The alpha-hidroxi-propionic acid utilization (compound I), linked to the 1,2,3-propanotriol or to the 1,2-propanodiol (compound II) in the sinusitis treatment, besides being a profitable alternative in the treatment of those diseases, will bring huge social and economic benefits to the country.

REQUESTS

1. "NEW UTILIZATION OF THE ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE": characterized by a pharmaceutic compound, consisting of the alpha-hidroxi-propionic acid (compound I), or an acceptable pharmaceutic solution of the latter, linked to 1,2,3-propanotriol, or to 1,2-propanodiol, or to serum or any other acceptable pharmaceutic vehicle, (compound II), having in the composition the dilution of: 0,2 to 0,5ml, or 1,1 to 1,9ml, or 2,0 to 3,0ml, or 3,1 to 4,0ml, or 4,1 to 5,0ml, or 5,1 to 6,0ml, or 6,1 to 10ml of compound I in 100ml of compound II; and still 0,3 to 0,8ml, or 0,4 to 0,7ml or 0,5 to 0,9ml, or 1,1 to 1,9 ml, or 2,0 to 2,5ml, or 2,5 to 3,0ml, or 3,1 to 4,0ml, or 4,1 to 5,0ml or 5,1 to 10,0ml of the active principle of compound I in 100ml of compound II.

2. "NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC IN MEDICINE", according to request I, characterized by its utilization in the sinusitis and other highairways diseases.

3. "NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC IN MEDICINE", according to request I, characterized by its utilization in the human and veterinarian highairway treatment.

4. "NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE", according to request I, characterized by its utilization as nasal releaser.

5. "NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE", according to request I, characterized by its dilution in 100ml of 1,2,3-propanotriol (compound II).

6. "NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE", according to request I, characterized by its utilization as a medication in the sinusitis and rhinitis treatment and as a clearing agent of the nasal cavities.

7. "NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC

09868793.100101

30 **ACID IN MEDICINE**", according to request I, characterized by its dilution in 100 ml of serum or any other acceptable pharmaceutic vehicle of the latter.

8. **"NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE"**, according to request I, characterized by its dilution in 100ml of serum or any other acceptable pharmaceutic vehicle of the latter.

35 9. **"NEW UTILIZATION OF ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE"**, according to request I, characterized by the dilution in 10ml of 1,2-propanodiol (compound II).

40 10. **"NEW UTILIZATION OF THE ALPHA-HIDROXI-PROPIONIC ACID IN MEDICINE"**, according to request I, characterized by being adapted for intake in drops, via nasal airways, or in the form of a solution for spraying, or a spray, a microfine powder for insufflation or an acceptable pharmaceutic salt of the latter, or an acceptable pharmaceutic solvate of the latter, so that a medicine can be prepared to the treatment of human and veterinarian highairways disturbances.

09868793.100101

add A1
add A2

add
C7

add
B2